

Phytochemical Analysis and Antibacterial Activities of *Combretum Molle* Stem Bark Extract for  
Management of Respiratory Tract Infections

BY

KAMPI MARIA GORRET

(BU/UP/2019/1536)

A Research project report Submitted to the Department of Chemistry for the Partial Fulfillment  
of the Requirements for the Award of the Bachelors of Science Education degree at Busitema  
University

**May 2023**

### **Declaration**

I, Kampi Maria Gorret, declare that the research dissertation is my own original work otherwise cited, and where such has been the case, references have been stated and that the same work has not been submitted for any award in any other university or other tertiary institute of higher education

Signature.....

Date... 24/05/23....

### **Approval**

This research review has been submitted for examination and has been approved by my supervisor.

Dr. Owor Richard Oriko

Signature..  Date... 24/05/23....

## **Dedication**

This report is dedicated to my parents and grandparents Ms. Kadondo Robinah and Mr. Mugabi Edward, my siblings Waiswa Micheal Mugabi, Kafuko Phillip, Mugabi David and Mpatogera Gertrude and my uncles Mr. Byakika Emmanuel Paddy, Mr. Kafuko Mosses, Mr. Ntabewra Hebert and Mr. Ibale Stephen Richard who have supported me in all aspects of life in my entire life journey of pursing my career dream as a professional teacher.

## **Acknowledgment**

I would like to express my sincere appreciation and gratitude to my supervisor Dr. Owor Richard Oriko for having guided me and supported me during each step of my research. My great appreciation also goes to the Busitema University Research and Innovation Fund (BURIF) for the financial support towards my work. My great thanks also go to the almighty God for the gift of life, the good health, protection and the wisdom entirely at school and throughout my life journey as a student. In great capacity I would like to thank the mighty fraternity of Busitema University, faculty of science and education, Nagongera campus which has been a great foundation in producing quality teachers with a sense of integrity.

With great value and respect to the department of Chemistry, and all the lecturers namely, Dr. Kamoga Omar, Dr. Oriko Richard Owor, Mr. Musagala Peter, Mr. Egor Moses, Dr. Andima Moses, Dr. Kigozi Moses, Mr. Katwesigye Richard and Ms. Nakijoba Lydia, for your entire support from the start of the course to the end.

I also thank my supervisor Dr. Oriko Richard Owor who always guided me and provided the necessary support towards the completion of my research review.

I would also thank prof. Olema David, Dr. Suubi Margret, Dr. Artubin, Mr. Kalyango, Mr. Wadaga Nelson, Dr. Owor Richard Oriko and Mr. Musagala Peter for having mentored me and guided as I was perusing my career.

Special regards go to Chemistry students more so Nambala Oliver, Wabwire Jacob, Kyotalye Brain, Kiwuso Hassan, Mwagale Flavia, Kisakye Kezia Priscilla and all the students of the faculty of science and education for the great company and support during my entire stay in Nagongera. Thanks for being good to me may God bless you all.

## **Abstract**

Respiratory tract infections affect both the upper and lower respiratory parts of the respiratory system. It's caused by the bacterium called *Streptococcus pneumoniae*. 9 million Ugandans are reported to have symptoms of respiratory tract infections after every two weeks in the primary health care centres and over 2 million people die each year. Traditionalists and herbalists in Uganda have adapted to using *Combretum molle* regulate the cases of respiratory tract infections. Therefore, this Study was to investigate the phytochemical composition of *C.molle* for antibacterial activities, analyze the phytochemical composition of *C.molle* and to evaluate the efficacy of *C.molle* stem back extract against antibacterial activities. The crude extract of *C.molle* was subjected to preliminary phytochemical screening and antimicrobial tests. The phytochemical tests were carried out using standard methods of analysis and these investigations revealed the presence of alkaloids, flavonoids, phenols, Quinones, tannins, Saponins and Glycosides while terpenoids and steroids were not present in the crude extract. An herbal syrup was formulated and named CODEM 40. This could top to greater protection and assistance to people in managing respiratory tract infections. Therefore, additional studies and elucidation of the active compounds so as to provide a new or principal component for production of new drugs can be opened up further studies to investigate the toxicity of *C.molle* herbal products before their recommendation for use.

## TABLE OF CONTENT

### Table of Contents

Declaration.....	i
Approval .....	i
Dedication.....	ii
Acknowledgment.....	iii
Abstract.....	iv
<b>TABLE OF CONTENT</b> .....	v
List of Acronyms .....	vii
List of figures.....	viii
List of tables.....	ix
<b>CHAPTER ONE: INTRODUCTION</b> .....	1
1.1 Background .....	1
1.2 Problem statement .....	3
1.3     Objectives.....	4
1.3.1 General objectives .....	4
1.3.2 Specific objectives .....	4
1.4 Significance .....	4
<b>CHAPTER TWO: LITERATURE REVIEW</b> .....	5
1.1 Description of a medicinal plant.....	5
2.2 <i>Combretum molle</i> .....	5
2.2.1 Structures of <i>Combretum molle</i> .....	7
2.2.2 PHTOCHEMICAL SCREENING OF <i>COMBRETUM MOLLE</i> .....	7
2.3 Respiratory Tract Infections.....	8
2.4 <i>Streptococcus pneumoniae</i> .....	8
2.4.1 STRUCTURE OF <i>STREPTOCOCCUS PNEUMONIAE</i> .....	9
2.5 Mode of transmission of Respiratory tract infections .....	9
2.6 Mechanisms Bacterial resistance against prescribed drugs.....	9
2.7 Diagnosis of Respiratory tract infections.....	10
2.8Treatment of respiratory tract infections .....	10
<b>CHAPTER THREE: MATERIAL AND METHODOLOGY</b> .....	11
3.1 Plant material .....	11

<b>3.2 Extraction .....</b>	11
<b>3.2.1 Aqueous extract (AE).....</b>	11
<b>3.2.2 Organic extract (OE) .....</b>	11
<b>3.3 Preliminary Phytochemical analysis; .....</b>	11
<b>3.3.1 Test for Alkaloids .....</b>	11
<b>3.3.2 Test for Flavonoids: .....</b>	12
<b>3.3.3 Tests for Phenols: .....</b>	14
<b>3.3.4 Test for Quinones:.....</b>	14
<b>3.3.5 Test for Tannins:.....</b>	15
<b>3.3.6 Test for Saponins: .....</b>	16
<b>3.4 Total flavonoid content: .....</b>	16
<b>3.5 Total alkaloid content:.....</b>	17
<b>3.6 Total phenolic content: .....</b>	18
<b>3.7 Total tannins.....</b>	19
<b>3.9 Formulation of a <i>Combretum molle</i> poly-herbal syrup.....</b>	19
<b>3.9.1 post formulation properties of CODEM 40 .....</b>	20
<b>3.9.2 PH test .....</b>	20
<b>3.9.3 Density determination .....</b>	20
<b>CHAPTER FOUR: RESULTS AND DISSCUSION .....</b>	21
<b>4.1 Phytochemical screening .....</b>	21
<b>4.2 properties of CODEM 40 .....</b>	23
<b>CHAPTER FIVE: CONCLUSION AND RECCOMENDATION .....</b>	25
<b>5.1 CONCLUSION .....</b>	25
<b>5.2 RECOMMENDATIONS.....</b>	25
<b>REFERENCES.....</b>	26
<b>Appendix.....</b>	33

## **List of Acronyms**

*C.molle; Combretum Molle*

*S. pneumoniae: Streptococcus pneumoniae*

WHO; world health organization

CMS; Carboxymethyl Cellulose

RTIs; Respiratory Tract Infections

HCl; Hydrochloric acid

AQ; Aqueous extract

OE; Organic extract

DNA; Deoxyribose Nucleic Acid

## List of figures

Figure 1 Structures of <b>Combretum molle</b> .....	7
Figure 2 structure of <b>streptococcus pneumoniae</b> .....	9
Figure 3 Wagner's Test for Alkaloids of C.molle .....	12
Figure 4 Dragendorff's reagent Test for alkaloids.....	12
Figure 5 <b>Ferric chloride test for Flavonoids</b> .....	13
Figure 6 <b>Zinc-HCl reduction test for Flavonoids</b> .....	13
Figure 7 <b>Lead-Acetate test for Flavonoids</b> .....	14
Figure 8 <b>ferric chloride test for phenols</b> .....	14
Figure 9 <b>Test for Quinones:</b> .....	15
Figure 10; <b>Ferric Chloride test for Tannins</b> .....	15
Figure 11 <b>Lead acetate test for tannins</b> .....	16
Figure 12 test for Saponins .....	16
Figure 13 A calibration curve for total flavonoid using standard quercetin solution. ....	17
Figure 14 A calibration curve for determination of total phenolic content of RT using standard ascorbic acid.....	18
Figure 15 A calibration curve for determination of total tannins using standard ascorbic acid ...	19
Figure 16 <b>A graph showing the phytochemical content</b> .....	23

## **List of tables**

<b>Table 1 THE PHYSIOCHEMICAL EVALUATION OF SYRUP FORMULATED .....</b>	<b>20</b>
<b>Table 2 phytochemical screening of the extracts of C. molle.....</b>	<b>21</b>
<b>Table 3 properties of CODEM 40 .....</b>	<b>23</b>

## **CHAPTER ONE: INTRODUCTION**

### **1.1 Background**

Respiratory tract infections (RTIs) are diseases associated with fever, sore throat, blocked nose, cough, colds, malaise and running nose. It is always caused either viruses like *Rhinovirus* and *Corona* virus or bacteria like *Streptococcus*, *Staphylococcus* and *Moraxella* (Chrian, Erasto, & Otieno, 2011). Globally, *Streptococcus* and *Staphylococcus* are the commonest bacteria that cause the majority of respiratory tract infections (Wiemken, Peyrani, & Ramirez, 2012).

A respiratory tract infection is one of the diseases that report the biggest number of patients in primary health care centers of Uganda. The commonest that affects the populations include pneumonia, asthma, and tuberculosis. These mainly affect extreme ages that is to say very young and very old (Wiemken et al., 2012). In Uganda, over 9 million people are infected by respiratory tract infections with about 2 million deaths every year. The majority of these infections occur in low income countries where the increased cases of infections are attributed to inaccessibility to immunization, medication and inability of health care systems to provide care thus leading to avoidable death (Boloursaz et al., 2013). In Uganda, 3 million infants (< 5 years) have been identified to be victims of respiratory tract infections every year (Kiguli et al., 2021). Entirely about 9 million Ugandans are reported to have symptoms of RTIs after every two weeks in the primary health care centres and over 2 million people die each year (Erku & Aberra, 2018).

There are numerous drugs prescribed for management of respiratory tract infections and they are in category of analgesics (paracetamol and ibuprofen), antihistamine (diphenhydramine), antibiotics (penicillin and amoxicillin) and anticholinergics (ipratropium and scopolamine) (WHO, 2001). In spite of the present drugs, RTIs cases are still in increase (Schuetz et al., 2013) For example in Uganda, there were 19 millions Ugandans suffering from respiratory tract infections in 2020, 18millions in 2021, 20millions in 2022. This is has contributed to COVID-19 (Vihta et al., 2022)

The surge up of the many victims suffering from RTIs is linked to resistance of the pathogens the available drugs (Cimolai, 2021).The cumulative rates of antibiotic resistance in community pathogens have focused the attention of researchers and clinicians on this public health problem(Ferri, Ranucci, Romagnoli, & Giaccone, 2017). There has been resistance to penicillin in

## REFERENCES

- Aberdein, J., Cole, J., Bewley, M., Marriott, H., & Dockrell, D. (2013). Alveolar macrophages in pulmonary host defence—the unrecognized role of apoptosis as a mechanism of intracellular bacterial killing. *Clinical & Experimental Immunology*, 174(2), 193-202.
- Ally, M. H. S. Antimicrobial Properties of Velvet Bush Willow (*Combretum molle*) Crude Bark Extracts on Selected Bacteria Species.
- Baggett, H. C., Watson, N. L., Deloria Knoll, M., Brooks, W. A., Feikin, D. R., Hammitt, L. L., . . . Madhi, S. A. (2017). Density of upper respiratory colonization with *Streptococcus pneumoniae* and its role in the diagnosis of pneumococcal pneumonia among children aged< 5 years in the PERCH study. *Clinical Infectious Diseases*, 64(suppl\_3), S317-S327.
- Beceiro, A., Tomás, M., & Bou, G. (2013). Antimicrobial resistance and virulence: a successful or deleterious association in the bacterial world? *Clinical microbiology reviews*, 26(2), 185-230.
- Boloursaz, M. R., Lotfian, F., Aghahosseini, F., Cheraghvandi, A., Khalilzadeh, S., Farjah, A., & Boloursaz, M. (2013). Epidemiology of lower respiratory tract infections in children. *Journal of Comprehensive Pediatrics*, 4(2), 93-98.
- Brooks, L. R., & Mias, G. I. (2018). *Streptococcus pneumoniae*'s virulence and host immunity: aging, diagnostics, and prevention. *Frontiers in immunology*, 9, 1366.
- Brusselaers, N., Vogelaers, D., & Blot, S. (2011). The rising problem of antimicrobial resistance in the intensive care unit. *Annals of intensive care*, 1, 1-7.
- Chrian, M., Erasto, P., & Otieno, N. J. (2011). Antimycobacterial activity and cytotoxicity effect of extracts of *Hallea rubrostipulata* and *Zanthoxylum chalybeum*. *Spatula DD*, 1(3), 147-152.
- Cimolai, N. (2021). The complexity of co-infections in the era of COVID-19. *SN Comprehensive Clinical Medicine*, 3(7), 1502-1514.
- Cock, I. (2015). The medicinal properties and phytochemistry of plants of the genus Terminalia (Combretaceae). *Inflammopharmacology*, 23, 203-229.
- Cock, I. E., & Van Vuuren, S. F. (2020). The traditional use of southern African medicinal plants for the treatment of bacterial respiratory diseases: A review of the ethnobotany and scientific evaluations. *Journal of Ethnopharmacology*, 263, 113204.

- Cummings, M. J., Tokarz, R., Bakamutumaho, B., Kayiwa, J., Byaruhangha, T., Owor, N., . . . Lutwama, J. J. (2019). Precision surveillance for viral respiratory pathogens: virome capture sequencing for the detection and genomic characterization of severe acute respiratory infection in Uganda. *Clinical Infectious Diseases*, 68(7), 1118-1125.
- Das, K., Tiwari, R., & Srivastava, D. (2010). Techniques for evaluation of medicinal plant products as antimicrobial agent: Current methods and future trends. *Journal of medicinal plants research*, 4(2), 104-111.
- de Menezes, B. R. C., Rodrigues, K. F., Schatkoski, V. M., Pereira, R. M., Ribas, R. G., do Amaral Montanheiro, T. L., & Thim, G. P. (2021). Current advances in drug delivery of nanoparticles for respiratory disease treatment. *Journal of Materials Chemistry B*, 9(7), 1745-1761.
- Dias, L. D., & Bagnato, V. S. (2020). An update on clinical photodynamic therapy for fighting respiratory tract infections: a promising tool against COVID-19 and its co-infections. *Laser Physics Letters*, 17(8), 083001.
- Eloff, J., Katerere, D., & McGaw, L. (2008). The biological activity and chemistry of the southern African Combretaceae. *Journal of Ethnopharmacology*, 119(3), 686-699.
- Erku, D. A., & Aberra, S. Y. (2018). Non-prescribed sale of antibiotics for acute childhood diarrhea and upper respiratory tract infection in community pharmacies: a 2 phase mixed-methods study. *Antimicrobial Resistance & Infection Control*, 7(1), 1-7.
- Ferri, M., Ranucci, E., Romagnoli, P., & Giaccone, V. (2017). Antimicrobial resistance: A global emerging threat to public health systems. *Critical reviews in food science and nutrition*, 57(13), 2857-2876.
- Fesseha, H., Kahsey, R., & Kidanemariam, F. (2019). An Insight Review on Antibiotic Resistance and Its Challenges. *Ind. J. Pure App. Biosci*, 7(6), 19-28.
- Friedman, N. D., Temkin, E., & Carmeli, Y. (2016). The negative impact of antibiotic resistance. *Clinical Microbiology and Infection*, 22(5), 416-422.
- Gabrielian, E., Shukarian, A., Goukasova, G., Chandanian, G., Panossian, A. G., Wikman, G., & Wagner, H. (2002). A double blind, placebo-controlled study of Andrographis paniculata fixed combination Kan Jang in the treatment of acute upper respiratory tract infections including sinusitis. *Phytomedicine*, 9(7), 589-597.

- Ghosh, B., & Mandal, S. (2018). Perspectives to DENV and GAS infections and associated serum LDH profiles. *EC Microbiology*, 14.
- Hassantabar, S., Ahmadi, M., & Sharifi, A. (2020). Diagnosis and detection of infected tissue of COVID-19 patients based on lung X-ray image using convolutional neural network approaches. *Chaos, Solitons & Fractals*, 140, 110170.
- Houghton, P., & Raman, A. (2012). *Laboratory handbook for the fractionation of natural extracts*: Springer Science & Business Media.
- Jin, Y.-H., Cai, L., Cheng, Z.-S., Cheng, H., Deng, T., Fan, Y.-P., . . . Huang, Q. (2020). A rapid advice guideline for the diagnosis and treatment of 2019 novel coronavirus (2019-nCoV) infected pneumonia (standard version). *Military medical research*, 7(1), 1-23.
- Kaggwa, B., Kyeyune, H., Munanura, E. I., Anywar, G., Lutoti, S., Aber, J., . . . Kamba, P. F. (2022). Safety and Efficacy of Medicinal Plants Used to Manufacture Herbal Products with Regulatory Approval in Uganda: A Cross-Sectional Study. *Evidence-based complementary and alternative medicine*, 2022.
- Kiguli, S., Olopot-Olupot, P., Alaroker, F., Engoru, C., Opoka, R. O., Tagoola, A., . . . Mogaka, C. (2021). Children's Oxygen Administration Strategies And Nutrition Trial (COAST-Nutrition): a protocol for a phase II randomised controlled trial. *Wellcome open research*, 6.
- Koevi, K.-K. A., Millogo, V., Fokou, J. B. H., Sarr, A., Ouedraogo, G. A., & Bassene, E. (2015). Phytochemical analysis and antioxidant activities of Combretum molle and Pericopsis laxiflora. *International Journal of Biological and Chemical Sciences*, 9(5), 2423-2431.
- Kulawe, D., Abubakar, A., & Abubakar, Z. (2019). The activities of the aqueous crude extracts of the leaf, root and bark of Combretum molle against selected test organisms. *International Journal of Scientific and Research Publications (IJSRP)*, 9(10), p9428.
- Lamorde, M., Tabuti, J. R., Obua, C., Kukunda-Byobona, C., Lanyero, H., Byakika-Kibwika, P., . . . Ryan, M. (2010). Medicinal plants used by traditional medicine practitioners for the treatment of HIV/AIDS and related conditions in Uganda. *Journal of ethnopharmacology*, 130(1), 43-53.
- Lappin, M., Blondeau, J., Boothe, D., Breitschwerdt, E., Guardabassi, L., Lloyd, D., . . . Turnidge, J. (2017). Antimicrobial use guidelines for treatment of respiratory tract disease in dogs and cats: Antimicrobial Guidelines Working Group of the International Society for

- Companion Animal Infectious Diseases. *Journal of Veterinary Internal Medicine*, 31(2), 279-294.
- Lewis, K. (2013). Platforms for antibiotic discovery. *Nature reviews Drug discovery*, 12(5), 371-387.
- Mahomoodally, M. F. (2013). Traditional medicines in Africa: an appraisal of ten potent African medicinal plants. *Evidence-based complementary and alternative medicine*, 2013.
- Majeed, M., & Rehman, R. U. (2021). Phytochemistry, Pharmacology, and Toxicity of an Epiphytic Medicinal Shrub *Viscum album* L.(White Berry Mistletoe). *Medicinal and Aromatic Plants: Healthcare and Industrial Applications*, 287-301.
- Martinez, J. L. (2014). General principles of antibiotic resistance in bacteria. *Drug Discovery Today: Technologies*, 11, 33-39.
- McGaw, L., Rabe, T., Sparg, S., Jäger, A., Eloff, J., & Van Staden, J. (2001). An investigation on the biological activity of *Combretum* species. *Journal of Ethnopharmacology*, 75(1), 45-50.
- Michael-Kordatou, I., Karaolia, P., & Fatta-Kassinos, D. (2018). The role of operating parameters and oxidative damage mechanisms of advanced chemical oxidation processes in the combat against antibiotic-resistant bacteria and resistance genes present in urban wastewater. *Water research*, 129, 208-230.
- Miedzybrodzki, R., Hoyle, N., Zhvaniya, F., Łusiak-Szelachowska, M., Weber-Dąbrowska, B., Łobocka, M., . . . Górska, A. (2021). Current updates from the long-standing phage research centers in Georgia, Poland, and Russia. *Bacteriophages: Biology, Technology, Therapy*, 921-951.
- Nadeem, S. F., Gohar, U. F., Tahir, S. F., Mukhtar, H., Pornpukdeewattana, S., Nukthamna, P., . . . Massa, S. (2020). Antimicrobial resistance: more than 70 years of war between humans and bacteria. *Critical Reviews in Microbiology*, 46(5), 578-599.
- Ncube, N., Afolayan, A., & Okoh, A. (2008). Assessment techniques of antimicrobial properties of natural compounds of plant origin: current methods and future trends. *African journal of biotechnology*, 7(12).
- Ntshanka M. N, E. I. P., Mthunzi F. M, Moloto M. J, Mubiayi K. P. Investigation into the Phytochemical Profile, Antioxidant and Antibacterial Potentials of *Combretum Molle* and *Acacia Mearnsii* Leaf Parts. *Biomed Pharmacol J* 2020;13(4). (2020).

- Nyenje, M. E. (2011). *Phytochemical Analysis And Bioactivity Of The Stem Bark Of Combretum Molle On Some Selected Bacterial Pathogens*. University of Fort Hare,
- O'Neill, J. (2016). Tackling drug-resistant infections globally: final report and recommendations.
- Organization, W. H. (2000). Overcoming antimicrobial resistance. *Overcoming antimicrobial resistance*.
- Pan, S.-Y., Zhou, S.-F., Gao, S.-H., Yu, Z.-L., Zhang, S.-F., Tang, M.-K., . . . Fong, W.-F. (2013). New perspectives on how to discover drugs from herbal medicines: CAM's outstanding contribution to modern therapeutics. *Evidence-based complementary and alternative medicine*, 2013.
- Papo, L., Van Vuuren, S., & Moteetee, A. (2022). The ethnobotany and antimicrobial activity of selected medicinal plants from Ga-Mashashane, Limpopo Province, South Africa. *South African Journal of Botany*, 149, 196-210.
- Planta, M. B. (2007). The role of poverty in antimicrobial resistance. *The Journal of the American Board of Family Medicine*, 20(6), 533-539.
- Raluca, P., & Mariana, B. DIFFERENTIAL DIAGNOSIS BETWEEN STAPHYLOCOCCI AND STREPTOCOCCI BY THE LATEX AGGLUTINATION TEST.
- Rammelkamp, C. H., & Maxon, T. (1942). Resistance of *Staphylococcus aureus* to the Action of Penicillin. *Proceedings of the Society for Experimental Biology and Medicine*, 51(3), 386-389.
- Reinert, R. (2009). The antimicrobial resistance profile of *Streptococcus pneumoniae*. *Clinical Microbiology and Infection*, 15, 7-11.
- Robertson, N. M., Nagourney, E. M., Pollard, S. L., Siddharthan, T., Kalyesubula, R., Surkan, P. J., . . . Kirenga, B. J. (2019). Urban-rural disparities in chronic obstructive pulmonary disease management and access in Uganda. *Chronic Obstructive Pulmonary Diseases: Journal of the COPD Foundation*, 6(1), 17.
- Roy, S., Gorai, D., Acharya, R., & Roy, R. (2014). *Combretum (combretaceae)*: Biological activity and phytochemistry. *American Journal of Pharm Research*, 4(11).
- Sandelowsky, H., Ställberg, B., Nager, A., & Hasselström, J. (2011). The prevalence of undiagnosed chronic obstructive pulmonary disease in a primary care population with respiratory tract infections-a case finding study. *BMC family practice*, 12(1), 1-9.

- Sarwar, A., & Imran, M. (2021). Prioritizing infection prevention and control activities for SARS-CoV-2 (COVID-19): a multi-criteria decision-analysis method. *Journal of healthcare leadership*, 77-84.
- Schuetz, P., Muller, B., Christ-Crain, M., Stolz, D., Tamm, M., Bouadma, L., . . . Tubach, F. (2013). Procalcitonin to initiate or discontinue antibiotics in acute respiratory tract infections. *Evidence-Based Child Health: A Cochrane Review Journal*, 8(4), 1297-1371.
- Schultz, F., Anywar, G., Wack, B., Quave, C. L., & Garbe, L.-A. (2020). Ethnobotanical study of selected medicinal plants traditionally used in the rural Greater Mpigi region of Uganda. *Journal of Ethnopharmacology*, 256, 112742.
- Sen, S., & Chakraborty, R. (2017). Revival, modernization and integration of Indian traditional herbal medicine in clinical practice: Importance, challenges and future. *Journal of traditional and complementary medicine*, 7(2), 234-244.
- Seukep, J. A., Sandjo, L. P., Ngadjui, B. T., & Kuete, V. (2016). Antibacterial and antibiotic-resistance modifying activity of the extracts and compounds from Nauclea pobeguinii against Gram-negative multi-drug resistant phenotypes. *BMC Complementary and Alternative Medicine*, 16(1), 1-8.
- Shakya, A. K. (2016). Medicinal plants: Future source of new drugs. *International journal of herbal medicine*, 4(4), 59-64.
- Shorrocks, B., & Bates, W. (2015). *The biology of African savannahs*: Oxford University Press, USA.
- Snidal, S. J., Barnard, G., Atuhairwe, E., & Amor, Y. B. (2015). Use of eCompliance, an innovative biometric system for monitoring of tuberculosis treatment in rural Uganda. *The American journal of tropical medicine and hygiene*, 92(6), 1271.
- Spellerberg, B., & Brandt, C. (2022). Laboratory diagnosis of Streptococcus pyogenes (group A streptococci).
- Ssenku, J. E., Okurut, S. A., Namuli, A., Kudamba, A., Tugume, P., Matovu, P., . . . Walusansa, A. (2022). Medicinal plant use, conservation, and the associated traditional knowledge in rural communities in Eastern Uganda. *Tropical Medicine and Health*, 50(1), 39.
- Tugume, P., & Nyakoojo, C. (2019). Ethno-pharmacological survey of herbal remedies used in the treatment of paediatric diseases in Buhunga parish, Rukungiri District, Uganda. *BMC Complementary and Alternative Medicine*, 19(1), 1-10.

- Umuhoza, T., Bulimo, W. D., Oyugi, J., Musabyimana, J. P., Kinengyere, A. A., & Mancuso, J. D. (2021). Prevalence of human respiratory syncytial virus, parainfluenza and adenoviruses in East Africa Community partner states of Kenya, Tanzania, and Uganda: A systematic review and meta-analysis (2007–2020). *PloS one*, 16(4), e0249992.
- Ventola, C. L. (2015). The antibiotic resistance crisis: part 1: causes and threats. *Pharmacy and therapeutics*, 40(4), 277.
- Vihta, K. D., Pouwels, K. B., Peto, T. E., Pritchard, E., Eyre, D. W., House, T., . . . Cook, D. (2022). Symptoms and Severe Acute Respiratory Syndrome Coronavirus 2 (SARS-CoV-2) Positivity in the General Population in the United Kingdom. *Clinical Infectious Diseases*, 75(1), e329-e337.
- Wang, L., Hu, C., & Shao, L. (2017). The antimicrobial activity of nanoparticles: present situation and prospects for the future. *International journal of nanomedicine*, 12, 1227.
- WHO. (2001). *Cough and cold remedies for the treatment of acute respiratory infections in young children*. Retrieved from
- Wiemken, T. L., Peyrani, P., & Ramirez, J. A. (2012). Global Changes in the Epidemiology of Community-Acquired Pneumonia. *Semin Respir Crit Care Med*, 33(03), 213-219.
- Woodhead, M., Blasi, F., Ewig, S., Huchon, G., Leven, M., Ortqvist, A., . . . Verheij, T. J. (2005). Guidelines for the management of adult lower respiratory tract infections. *European Respiratory Journal*, 26(6), 1138-1180.
- Zoorob, R., Sidani, M. A., Fremont, R. D., & Kihlberg, C. (2012). Antibiotic use in acute upper respiratory tract infections. *American family physician*, 86(9), 817-822.